

National Imaging Associates, Inc.*	
Clinical guidelines	Original Date: October 2009
MYOCARDIAL PERFUSION IMAGING	
(aka NUCLEAR CARDIAC IMAGING STUDY)	
CPT Codes: 78451, 78452, 78453, 78454, 78466,	Last Revised Date: March-February 20221
78468, 78469, 78481, 78483, 78499	
Guideline Number: NIA_CG_024	Implementation Date: January 20232

GENERAL INFORMATION

It is an expectation that all patients receive care/services from a licensed clinician. All appropriate supporting documentation, including recent pertinent office visit notes, laboratory data, and results of any special testing must be provided. All prior relevant imaging results, and the reason that alternative imaging cannot be performed, must be included in the documentation submitted.

This guideline is for stress imaging, specifically myocardial perfusion imaging (MPI), with appropriate preference for suitable alternatives, such as stress echocardiography (SE), when more suitable, unless otherwise stated (refer to Overview section).

INDICATIONS for MPI¹⁻⁴

(Fihn 2012, Hendel 2009, Montalescot 2013, Wolk 2014)

SUSPECTED Coronary Artery Disease CORONARY ARTERY DISEASE (CAD)

- Symptomatic patients without known CAD (use <u>Diamond Forrester Table</u>)
 - Low or intermediate pretest probability and unable to exercise exercise (SE diversion not required)

- High pretest probability (<u>SE diversion not required</u>) (Stress Echocardiogram [SE] diversion not required)
- Repeat testing in a patient with new or worsening symptoms and negative result at least one year prior AND meets one of the criteria above

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^{1—} Myocardial Perfusion Imaging

Asymptomatic patients without known CAD (SE diversion not required)

- Previously unevaluated ECG evidence of possible myocardial ischemia including ischemic
 ST segment or T wave abnormalities (See <u>Overview section</u>)
- Previously unevaluated pathologic Q waves <u>(see section in Overview)(see Overview</u> section)
- Previously unevaluated complete left bundle branch block
- Those with History of diabetes mellitus, > 40 years olda, with calcium score >400⁵, not previously evaluated⁶ (Budoff, 2016)

ABNORMAL CALCIUM SCORES (CAC)⁴⁻⁸

- ASYMPTOMATIC patient with a calcium score >400, not previously evaluated
- SYMPTOMATIC patient with prior CAC ≥100

INCONCLUSIVE CAD EVALUATION WITHIN THE PAST 2 YEARS AND OBSTRUCTIVE CAD REMAINS A CONCERN

- Exercise stress ECG with low-risk Duke treadmill score (≥5), (see section in Overview)e
 Overview) but patient's current symptoms indicate an intermediate or high pretest probability
 (SE diversion not required (SE diversion not required for high pretest probability)
- Exercise stress ECG with an intermediate Duke treadmill score
- Intermediate coronary computed tomography angiography (CCTA) (e.g.e.g., 430 70% lesions)
- Non-diagnostic exercise stress test with inability to achieve target heart rate (THR) (SE diversion not required)
- An indeterminate (equivocal, borderline, or discordant) evaluation by prior stress imaging (SE or CMR) within the past 2 years (SE diversion not required)

FOLLOW-UP OF PATIENT'S POST CORONARY REVASCULARIZATION (PCI or CABG)4 (Wolk, 2014)

Asymptomatic follow-up stress imaging (MPI or SE) at a minimum of 2 years post coronary artery bypass grafting (CABG) or percutaneous coronary intervention (PCI) (whichever is later) is appropriate for patients with a history of silent ischemia or a history of a prior left main stent. (SE diversion not required for CABG) (Wolk, 2014)

OR

For patients with high occupational risk-(e.g., associated with public safety, airline and boat pilots, bus and train drivers, bridge and tunnel workers/toll collectors, police officers and firefighters (SE diversion not required)

 New, recurrent, or worsening symptoms post coronary revascularization is an indication for stress imaging-(), if it will alter management

FOLLOW-UP OF KNOWN CAD

- Low or intermediate pretest probability chest pain with coronary disease defined by a Coronary calcium score ≥100 is ok for stress imaging
- Follow-up of asymptomatic or stable symptoms when last invasive or non-invasive assessment of coronary disease showed hemodynamically significant CAD (ischemia on stress test or FFR ≤ 0.80 or stenosis ≥ 70% of a major vessel), over two years ago, without intervening coronary revascularization is an appropriate indication for stress imaging (MPI or SE) in patients if it will alter management

SPECIAL DIAGNOSTIC CONDITIONS REQUIRING CORONARY EVALUATION

- Prior acute coronary syndrome (with documentation in MD notes), without invasive or non-invasive coronary evaluation (<u>SE diversion not required</u>)
- Newly diagnosed systolic heart failure (EF < 50%) or diastolic heart failure, with reasonable suspicion of cardiac ischemia (prior events, risk factors), with symptoms or signs of ischemia unless invasive coronary angiography is immediately planned (SE diversion not required)^{1, 9-11} (Fihn, 2012; Patel, 2013; Yancy, 2013) we hav
- LVEF ≤ 50%-requiring myocardial viability assessment to assist with decisions regarding coronary revascularization^{9, 10} (Patel, 2013; Yancy, 2013)
- Ventricular arrhythmias
 - Sustained ventricular tachycardia (VT) > 100 bpm, ventricular fibrillation (VF), or exercise-induced VT, when invasive coronary arteriography is not immediately planned¹² (Al Khatib 2018) (SE diversion not required)
 - Nonsustained VT, multiple episodes, each ≥ 3 beats at ≥ 100 bpm, or frequent PVCs (defined as greater than or equal to 30/hour on remote monitoring) without known cause or associated cardiac pathology, when an exercise ECG cannot be performed¹³ (Zimetbaum 2018)
- Prior to initiation of Class IC antiarrhythmic drug initiation (Propafenone or Flecainide), as well as annually in intermediate and high global risk patients (SE diversion not required) (SE diversion not required)¹⁴ (Reiffel, 2015)
- Assessment of hemodynamic significance of one of the following documented conditions:
 - Anomalous coronary arteries¹⁵ (Grani, 2017)
 - Myocardial bridging of coronary artery
- Coronary aneurysms in Kawasaki's disease¹⁶ (Newburger, 2016) or due to atherosclerosis
- Following radiation therapy to the anterior or left chest, at 5 years post initiation and every 5 years thereafter¹⁷ (Lancellotti, 2013)

PRIOR TO ELECTIVE NON-CARDIAC SURGERY IN ASYMPTOMATIC PATIENTS

- An ilntermediate or high risk surgery with of one or more risk factors (see below), AND documentation of an inability to walk (or <4 METs) AND there has not been an imaging stress test within 1 year¹⁸⁻²⁰*
 - Patients who have no above indication for non-invasive coronary evaluation, but are referred for preoperative cardiac evaluation, are eligible for MPI if all 4 criteria are met:
 - Surgery is supra-inguinal vascular, intrathoracic, or intra-abdominal; AND
 - On The patient has at least one of the additional cardiac complication risk factors:
 - → Ischemic Heart Disease
 - History of stroke or transient ischemic attack (TIA)
 - → History of congestive heart failure or ejection fraction ≤ 35%
 - Insulin-requiring diabetes mellitus

 - → AND
 - The patient has limited functional capacity (< 4 METS), such as one of the following:
 - Unable to take care of their activities of daily living (ADLs) or ambulate
 - Unable to walk 2 blocks on level ground
 - Unable to climb 1 flight of stairs
 - → AND
 - There has not been a conclusive stress evaluation, CTA, or heart catheterization within the past year; and the results of such a test would be likely to substantially alter therapy and/or preclude proceeding with the intended surgery.
 - Risk factors: history of ischemic heart disease, history of congestive heart failure, history of cerebrovascular disease, preoperative treatment with insulin, and preoperative serum creatinine >2.0 mg/dL.
 - Surgical Risk:
 - High risk surgery: Aortic and other major vascular surgery, <u>p</u>Peripheral vascular surgery, <u>a</u>Anticipated prolonged surgical procedures associated with large fluid shifts and/or blood loss
 - Intermediate risk surgery: Carotid endarterectomy, head and neck surgery, intraperitoneal and intrathoracic surgery, oorthopedic surgery, perostate surgery
 - —<u>Low risk surgery</u>: <u>Endoscopic procedures, superficial procedure, cataract surgery</u>, <u>bB</u>reast surgery
 - ___
- Planning for <u>anysolid</u> organ <u>or stem cell</u> transplantation is an indication for preoperative MPI, if there has not been a conclusive stress evaluation, CTA, or heart catheterization within the past year, <u>-at the discretion of the transplant service</u>. <u>-and with ≥ 3 of the following risk factors: (SE diversion not required)^{3, 21} (<u>Lentine</u>, <u>2012</u>):
 </u>

- Age > 60
- Smoking
- Hypertension
- Dyslipidemia
- Left ventricular hypertrophy
- > 1 year on dialysis (for renal transplant patients)
- Diabetes mellitus
- Prior ischemic heart disease

POST CARDIAC TRANSPLANT (SE diversion not required) (SE diversion not required)

- Annually, for the first five years post cardiac transplantation, in a patient not undergoing invasive coronary arteriography
- After the first five years post cardiac transplantation, patients with documented transplant coronary vasculopathy can be screened annually unless invasive coronary arteriography is planned

BACKGROUND

This guideline is for stress imaging, specifically myocardial perfusion imaging (MPI), with appropriate preference for alternatives, such as stress echocardiography (SE) or stress ECG alone when more suitable (see section below).

Radionuclide myocardial perfusion imaging (MPI) allows for evaluation of cardiac perfusion at rest and at exercise, as well as using pharmacologic agents for the diagnosis and management of coronary artery disease. With radionuclide MPI, pharmacologic stress may be performed with an inotropic agent or vasodilator. These agents are indicated for patients who cannot reach an adequate endpoint with physical exercise stress testing (Pagnanelli, 2017).²²

Stable patients without known CAD fall into 2 categories^{1, 3, 4} (Fihn, 2012; Montalescot, 2013; Wolk, 2013):

- Asymptomatic, for whom global risk of CAD events can be determined from coronary risk factors, using calculators available online (see <u>Websites for Global Cardiovascular Risk</u> <u>Calculators</u> section).
- **Symptomatic,** for whom we estimate the pretest probability that their chest-related symptoms are due to clinically significant CAD (below):

The 3 Types of Chest Pain or Discomfort

- Typical Angina (Definite) is defined as including all 3 characteristics:
 - Substernal chest pain or discomfort with characteristic quality and duration
 - Provoked by exertion or emotional stress

- o Relieved by rest and/or nitroglycerine
- Atypical Angina (Probable) has only 2 of the above characteristics
- Nonanginal Chest Pain/Discomfort has only 0 1 of the above characteristics

<u>The medical record should provide enough detail to establish Once</u> the type of chest pain. <u>From those details, has been established from the medical record, t</u>The Pretest Probability of obstructive CAD is estimated from the **Diamond Forrester Table** below, recognizing that in some cases multiple additional coronary risk factors could increase pretest probability^{1, 4} (Fihn 2012, Wolk 2013):

Diamond Forrester Table

Age (Years)	Gender	Typical/Definite Angina Pectoris	Atypical/Probable Angina Pectoris	Nonanginal Chest Pain
< 20	Men	Intermediate	Intermediate	Low
≤ 39	Women	Intermediate	Very low	Very low
40.40	Men	High	Intermediate	Intermediate
40–49	Women	Intermediate	Low	Very low
FO FO	Men	High	Intermediate	Intermediate
50–59	Women	Intermediate	Intermediate	Low
> 60	Men	High	Intermediate	Intermediate
≥ 60	Women	High	Intermediate	Intermediate

- Very low: < 5%pretest probability of CAD, usually not requiring stress evaluation
- Low: 5 10% pretest probability of CAD
- Intermediate: 10% 90% pretest probability of CAD
- High: > 90% pretest probability of CAD

OVERVIEW

MPI may be performed without diversion to <u>a_SE</u> in any of the following^{4, 23} (Henzlova, 2016; Wolk, 2013):

- Inability to Exercise
 - Physical limitations precluding ability to exercise for at least 3 full minutes of Bruce protocol
 - → BMI > 40kg/m²
 - Limited functional capacity (< 4 METS) such as one of the following:
 - Unable to take care of their ADLs or ambulate
 - Unable to walk 2 blocks on level ground
 - Unable to climb 1 flight of stairs
- Other Comorbidities
 - Severe chronic obstructive pulmonary disease (COPD) with pulmonary function test (PFT) documentation, severe shortness of breath on minimal exertion, or requirement of home oxygen during the day
 - Poorly controlled hypertension, with systolic BP > 180 or diastolic BP > 120 (and clinical urgency not to delay MPI)

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- ECG and Echo-Related Baseline Findings
 - Prior cardiac surgery (coronary artery bypass graft or valvular)

- Documented poor acoustic imaging window
- Left ventricular ejection fraction ≤ 40%
- Pacemaker or ICD
- Persistent atrial fibrillation
- Resting wall motion abnormalities that would make SE interpretation difficult
- Complete left bundle branch block (LBBB)

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- Risk-Related scenarios
 - High pretest probability in suspected CAD
 - Intermediate or high global risk in patients requiring type IC antiarrhythmic drugs (prior to initiation of therapy and annually)
 - Arrhythmia risk with exercise

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- Previously unevaluated pathologic Q waves (in two contiguous leads) defined as the following:
 - **e**→> 40 ms (1 mm) wide

 - e-> 25% of depth of QRS complex
 ECG Stress Test Alone versus Stress Testing with Imaging

ECG Stress Test Alone versus Stress Testing with Imaging

Prominent scenarios suitable for an ECG stress test WITHOUT imaging (i.e., exercise treadmill ECG test) require that the patient can exercise for at least 3 minutes of Bruce protocol with achievement of near maximal heart rate **AND** has an interpretable ECG for ischemia during exercise⁴ (Wolk, 2014):

- The (symptomatic) low or intermediate pretest probability patient who can exercise and has an interpretable ECG⁴ (Wolk, 2014)
- The patient who is under evaluation for exercise-induced arrhythmia
- The patient who requires an entrance stress test ECG for a cardiac rehab program or for an exercise prescription
- For the evaluation of syncope or presyncope during exertion²⁴ (Shen, 2017)

Duke Exercise ECG Treadmill Score²⁵ (Mark. 1987)

Calculates risk from ECG treadmill alone:

- The equation for calculating the Duke treadmill score (DTS) is: DTS = exercise time in minutes (5 x ST deviation in mm or 0.1 mV increments) (4 x exercise angina score), with angina score
 being 0 = none, 1 = non-limiting, and 2 = exercise-limiting
- The score typically ranges from 25 to + 15. These values correspond to low-risk (with a score of ≥ + 5), intermediate risk (with scores ranging from 10 to + 4), and high-risk (with a score of ≤ 11) categories

An uninterpretable baseline ECG includes (Fihn, 2012):

- ST segment depression 1 mm or more; (not for non-specific ST- T wave changes)
- Ischemic looking T waves; at least 2.5 mm inversions (excluding V1 and V2)

- LVH with repolarization abnormalities, pre-excitation pattern such as WPW, ventricular paced rhythm, or LBBB
- Digitalis use with associated ST segment abnormalities
- Resting HR under 50 bpm on a medication, such as beta-blockers or calcium channel blockers, that is required for patient's treatment and cannot be stopped, with an anticipated suboptimal workload

Global Risk of Cardiovascular Disease

Global risk of CAD is defined as the probability of manifesting cardiovascular disease over the next 10 years and refers to **asymptomatic** patients without known cardiovascular disease. It should be determined using one of the risk calculators below. A high risk is considered greater than a 20% risk of a cardiovascular event over the ensuing 10 years. **High global risk by itself generally lacks scientific support as an indication for stress imaging.** There are rare exceptions, such as patients requiring IC antiarrhythmic drugs who might require coronary risk stratification prior to initiation of the drug-or patients with a CAC score > 400 Agastson units, when global risk is moderate or high.

- CAD Risk—Low
 - 10-year absolute coronary or cardiovascular risk less than 10%.
- CAD Risk—Moderate
 - 10-year absolute coronary or cardiovascular risk between 10% and 20%.
- CAD Risk—High
 - 10-year absolute coronary or cardiovascular risk of greater than 20%.

Websites for Global Cardiovascular Risk Calculators *26-30

(Arnett, 2019; D'Agostino, 2008; Goff, 2014; McClelland, 2015; Ridker, 2007)

*Patients who have already manifested cardiovascular disease are already at high global risk and are not applicable to the calculators.

Risk Calculator	Websites for Online Calculator
Framingham Cardiovascular	https://reference.medscape.com/calculator/framingham-
Risk	<u>cardiovascular-disease-risk</u>
Reynolds Risk Score	http://www.reynoldsriskscore.org/
Can use if no diabetes	
Unique for use of family	
history	
Pooled Cohort Equation	http://clincalc.com/Cardiology/ASCVD/PooledCohort.aspx?example
ACC/AHA Risk Calculator	http://tools.acc.org/ASCVD-Risk-Estimator/

MESA Risk Calculator	https://www.mesa-
With addition of Coronary	nhlbi.org/MESACHDRisk/MesaRiskScore/RiskScore.aspx
Artery Calcium Score, for CAD-	
only risk	

Definitions of Coronary Artery Disease^{1, 3, 6, 31}

(Fihn, 2012; Mintz, 2016; Montalescot, 2013; Patel, 2017)

Percentage stenosis refers to the reduction in diameter stenosis when angiography is the method and can be estimated or measured using angiography or more accurately measured with intravascular ultrasound (IVUS).

- Coronary artery calcification is a marker of risk, as measured by Agatston score on coronary artery calcium imaging. It is not a diagnostic tool so much as it is a risk stratification tool. Its incorporation into global risk can be achieved by using the MESA risk calculator.
- Ischemia-producing disease (also called hemodynamically or functionally significant disease, for which revascularization might be appropriate) generally implies at least one of the following:
 - Suggested by percentage diameter stenosis ≥ 70% by angiography; intermediate lesions are 50 69%³²borderline lesions are 40 70%⁴ (Fihn, 2012)
 - For a left main artery, suggested by a percentage stenosis ≥ 50%^{1, 31, 33} (Fihn, 2012; Lofti, 2018; Mintz, 2016)
 - FFR (fractional flow reserve) ≤ 0.80 for a major vessel^{31, 33} (Lofti, 2018; Mintz, 2016)
 - Demonstrable ischemic findings on stress testing (ECG or stress imaging), that are at least mild in degree
- FFR (fractional flow reserve) is the distal to proximal pressure ratio across a coronary lesion. Less than or equal to 0.80 is considered a significant reduction in coronary flow.

Anginal Equivalent^{1, 24}

(Fihn, 2012; Shen, 2017)

Development of an anginal equivalent (e.g., shortness of breath, fatigue, or weakness) either with or without prior coronary revascularization should be based upon the documentation of reasons to suspect that symptoms other than chest discomfort are not due to other organ systems (e.g., dyspnea due to lung disease, fatigue due to anemia). This may include respiratory rate, oximetry, lung exam, etc. (as well as d-dimer, chest CT(A), and/or PFTs, when appropriate), and then incorporated into the evaluation of coronary artery disease as would chest discomfort. Syncope per se is not an anginal equivalent.

- High risk surgery: Aortic and other major vascular surgery, Peripheral vascular surgery, Anticipated prolonged surgical procedures associated with large fluid shifts and/or blood loss
- Intermediate risk surgery: Carotid endarterectomy, Head and neek surgery, Intraperitoneal and intrathoracie surgery, Orthopedie surgery, Prostate surgery
- Low risk surgery: Endoscopic procedures, superficial procedure, cataract surgery, Breast surgery

Abbreviations

ADLs	Activities of daily living
BSA	Body surface area in square meters
CABG	Coronary artery bypass grafting
CAD	Coronary artery disease
CMR	Cardiac magnetic resonance imaging
CTA	Computed tomography angiography
ECG	Electrocardiogram
FFR	Fractional flow reserve
IVUS	Intravascular ultrasound
LBBB—LBBB	Left bundle-branch block
LVEF	Left ventricular ejection fraction
LVH	Left ventricular hypertrophy
MI	Myocardial infarction
MET	Estimated metabolic equivalent of exercise
MPI	Myocardial perfusion imaging
PCI	Percutaneous coronary intervention
PFT	Pulmonary function test
PVCs	Premature ventricular contractions
SE	Stress echocardiography
THR	Target heart rate
VT	Ventricular tachycardia
VF	Ventricular fibrillation
WPW	Wolf Parkinson White

POLICY HISTORY

<u>Date</u>	Summary
March February 2022	 BroughtMoved the sentence regarding utilization of suitable
	alternatives such as Stress Echocardiography to the General
	Information section
	 Placed Link to Overview Section in General Information
	 Clarified evaluation of possible ischemia in newly diagnosed heart
	failure by stating "with reasonable suspicion of cardiac ischemia
	(prior events, risk factors, or symptoms and signs)"
	 Clarified "intermediate lesions are 50-69%" for ischemia-producing
	<u>disease</u>
	 Added stress imaging approval for calcium score > 100 with low to
	intermediate probability symptoms
	 Deleted the requirement for diabetes when calcium score > 400 for
	stress imaging

- Deleted "≤50%" from "LVEF ≤50% requiring myocardial viability assessment to assist with decisions regarding coronary revascularization"
- Added Calcium score section:
 - Added stress imaging approval for calcium score > 100 with symptoms consistent with low to intermediate pretest probability
- Added Calcium score section
- Added SE diversion not required in section regarding 'An indeterminate (equivocal, borderline, or discordant) evaluation by prior stress imaging (SE or CMR) within the past 2 years'
- Added reminder (SE diversion not required for CABG)
- Changed preoperative guideline to include intermediate risk surgery
 with one or more risk factors AND documentation of an inability to
 walk (or <4 METs) AND there has not been an imaging stress test
 within 1 year
- Changed solid organ transplant guideline to include stem cell transplant and "any" organ transplant
- Added definition of surgical risk to preop guidelines Changed
 preoperative guideline to An Intermediate or high risk surgery with of
 one or more risk factors. AND documentation of an inability to walk
 (or <4 METs) AND there has not been an imaging stress test within 1
 year*
- Added risk factors for preop guidelines and definition of surgical risk to overview section
- Changed solid organ transplant guideline
- •
- In Background section clarified the requirement for description of chest pain by adding sentence "The medical record should provide enough detail to establish the type of chest pain."
- Added definition of Q waves
- Deleted sentence regarding calcium scoring within the Global Risk
 Section
- Deleted sentence regarding using calcium score solely for risk stratification
- Transplant stress testing modified
- Deleted IFR references

March 2021

- Wording changes for low and intermediate pretest probability patients
- Added annual studies for patients on Flecainide
- Added indication for Ca score in diabetic > 40 and calcium score > 400
 with reference added
- ----Removed BMI > 40 as indication for MPI
- •

Name and 2020	Addition of the state of the st
March 2020	Added general information section as Introduction which outlines
	requirements for documentation of pertinent office notes by a
	licensed clinician, and inclusion of laboratory testing and relevant
	imaging results for case review
	 Added clarification of repeat testing in a patient with new or
	worsening symptoms and negative result at least one year prior to
	include the statement "AND meets one of the criteria above"
	 Added clarification of frequent PVCs under ventricular arrhythmias
	which states defined as greater than or equal to 30/hour to include
	"on remote monitoring"
	• Edited indication of planning for solid organ transplantation to remove
	the requirement of limited functional capacity but maintaining
	requirement of ≥ 3 listed risk factors
	 Removed explanation of three vasodilators approved for stress testing
	from the background
	Added edits to the Coronary Artery disease coronary artery disease
	definition section
	Updated and added new references
	•
July 2019	For special diagnostic consideration, prior acute coronary syndrome
	(as documented in MD notes), the following clause was added:
	'without subsequent invasive or non-invasive coronary evaluation (SE
	diversion not required)'
	For section on prior to elective non-cardiac surgery the following was
	added: 'There has not been a conclusive stress evaluation, CTA, or
	heart catheterization within the past year'
	For section on prior to elective non-cardiac surgery indication
	'Planning for solid organ transplantation is an indication for
	preoperative MPI, if there has not been a conclusive stress evaluation,
	CTA, or heart catheterization within the past year'
	Added indication for follow-up every 2 years for patients with known
	CAD in high-risk occupations
	Added prior left main stent in asymptomatic patients as follow-up
	every two years
	Clarification of diversion to stress echo in suitable patients' post-
	revascularization
	Clarification of post cardiac transplant
	Removed section on Global Risk Calculator
	Added "with EKGECG changes," as indication for stress echo in patients an discovin or with LVH
	on digoxin or with LVH
	Removed indication for ETT in asymptomatic patients Added an appropriate and appropriate
	Added presyncope and syncope with exercise as an indication for ETT
	•

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